**Plastic Surgery** 



# MDR WITH METALLO-BETA-LACTAMASE-PRODUCING PSEUDOMONAS AERUGINOSA ISOLATED IN BURN PATIENTS AT A TERTIARY CARE TEACHING HOSPITAL

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ABSTRACT BACKGROUND: Metallo-beta-lactamase (MBL) delivering *Pseudomonas aeruginosa* in the consume patients is a main source of dreariness and mortality and remains a genuine wellbeing worry among the clinicians. OBJECTIVES: The point of this investigation was to recognize MBL-delivering *P. aeruginosa* in consume patients and decide multidrug-safe (MDR) strains, and individual opposition designs.

**PATIENTS AND METHODS:** In this cross-sectional examination, 270 strains of *P. aeruginosa* were secluded from the consume patients alluded to Department of plastic surgery, IMS and SUM hospital, Bhubaneswar. Among them, 55 MBL-delivering *P. aeruginosa* strains were confined from 55 patients hospitalized in consume unit. Least inhibitory fixations (MICs) and MBLs were dictated by the E-test strategy. **RESULTS:** Of the 55 consume cases, 29 (53%) were females and 26 (47%) guys. Harmed consume patients' ages ran from 16 to 87 years, with most extreme number of cases in the age gathering of 16 to 36 years (n, 40; 72.7%). By and large, 32 cases were coincidental (60%), and 22 were self-destructive consumes (40%). Of the 55 consume patients, 17 cases were lapsed (30%). All passings were because of concoction exposures. In anti-toxin defenselessness testing by E-test technique, ceftazidime was the best one and 35 disconnects (63.5%) were impervious to all the 11 tried anti-infection agents.

**CONCLUSIONS:** Routine microbiological observation and cautious in vitro testing of anti-infection agents before remedy and strict adherence to healing center anti-infection strategy may counteract, treat, and control MDR and pandrug - safe (PDR) *P. aeruginosa* strains in consume units.

KEYWORDS : Burn patients; Pseudomonas aeruginosa; Metallo-Beta-Lactamase; Drug Resistance

## INTRODUCTIONS

Pseudomonas aeruginosa (P. aeruginosa) is a main source of nosocomial diseases in consume patients. For treatment .of these contaminations, carbapenems, particularly imipenem, are utilized. Nonetheless, the commonness of imipenem protection from P. aeruginosa has been expanding around the world. Protection from carbapenems is because of impermeability by means of the loss of the OprD porin, the up-direction of a functioning efflux siphon arrangement of the cytoplasmic layer, or the generation of metallo-βlactamases (MBLs). The nearness of these components can ead to treatment disappointment in carbapenem treatment of P. aeruginosa infections.1 The MBL chemicals effectively hydrolyze all β-lactams, for example, penicillins, cephalosporins, and carbapenems, with the exception of aztreonam in vitro.2 These compounds have a place with Ambler class B and Bush gathering 3 and require divalent cations, normally zinc, as a cofactor for protein movement. They are hindered by metal chelators, for example, EDTA yet are not influenced by remedial β-lactamase inhibitors like sulbactam, tazobactam, or clavulanic acid.3 Metallo-\beta-lactam qualities are normally part of an integron structure and are carried on transferable plasmids yet can likewise be a piece of the chromosome. In light of the integronassociated quality tapes, MBLs-creating P. aeruginosa segregates are regularly impervious to various gatherings of antimicrobial specialists, which can be exchanged to different sorts of bacteria.4 Therefore, MBLs-delivering strains are critical from a disease control point of view. The MBLs are partitioned into the accompanying 6 bunches dependent on atomic structure: IMP, VIM, SIM, SPM, GIM, and AIM.5 Metallo-β-lactamases-creating P. aeruginosa segregates were first detailed in Japan in 1991, and from that point forward have been recognized in different countries.6 There are 2 reports about the seclusion of MBL-delivering P. aeruginosa from consume patients in southern Iran.7,8 The present investigation was finished to distinguish MBL-creating P. aeruginosa disengaged from patients admitted to IMS and SUM hospital, Bhubaneswar.

## **PATIENTS AND METHODS**

In this cross-sectional examination, 270 strains of P. aeruginosa were separated from the consume patients alluded to Ghot-beddin Burn Hospital, Shiraz, Iran, amid 2009-2010. Information concerning sexual orientation, age, length of hospitalization, cause, site, degree, and sorts of consumes (coincidental or sui-cidal) were gathered from the patients tainted with MBL-delivering P. aeruginosa through polls filled by gifted medical attendants. This examination was performed as per the moral benchmarks set down in the 1964 Helsinki presentation. Every one of the patients were relegated code num-bers. The plan and convention of the investigation were endorsed by the Ethics Committee of Professor Alborzi Clinical Microbiology Research Center (PACMRC), Shiraz, Iran.

#### Isolation and Identification of Metallo-Beta- Lactamase-Producing Pseudomonas aeruginosa

Amid a one-year time frame, 55 MBL-creating P. aerugi-nosa strains were disconnected from 55 copy patients hospital-ized in the copy unit. The detachment and ID of P. aeruginosa from wound examples were performed by the accompanying ordinary strategies. Examples were gathered by sterile swabs after the evacuation of dressing and purging the injury surface by 70% liquor. The consume tests were refined on supplement agar (NA; Oxoid Ltd, London, UK) and brooded at 35°C to 37°C medium-term. At that point any suspicious settlement was subcultured and refined. The detaches were recognized as P. aeruginosa, in light of Gram recoloring, catalase test, oxidase test, triple sugar press (TSI) maturation, motility, shading, pyocyanin color produc-tion, and scent. For definite affirmation, biochemical tests implanted in the API-20E biochemical unit framework (Bio-Mer-ieux, France) and manual biochemical tests were utilized as per the producer's guidelines. Strains were saved at - 20°C on tryptic soy soup (TSB; Oxoid Ltd, London, UK) containing 20% (v/v) glycerol. MBL E-test strips (AB BIODISK, Solna, Sweden) were utilized to screen class B beta-lactamase. Tests were performed and inter-preted as per the producer's guidelines.

### Antibiotic Susceptibility And Resistance Patterns

Least inhibitory focuses (MICs) of the 11 antimicrobial specialists routinely recommended in consume cen-ters against the 55 confines of MBL-creating P. aeruginosa were dictated by the E-test technique (AB BIODISK, Solna, Sweden), as prescribed by the National Com-mittee For Clinical Laboratory Standard Institute (CLSI) (19). A bacterial suspension from development in a tryptic soy agar (TSA) plate was set up in 2 mL of Mueller-Hinton juices (MHB), and the turbidity was balanced with the goal that it was equal to that of a 0.5 McFarland standard. The bacterial suspension was streaked onto a 150-mm-diam-eter plate containing Mueller-Hinton agar (MHA); the plate was later hatched at 35 to 37°C in encompassing air for 16 to 18 hours. The MIC was perused based on the inter-ception of the curved zone of development hindrance with the evaluated E-test strip as indicated by the maker's directions. The anti-infection agents (Mast Co., UK) comprised of imipenem (10 µg), meropenem (10 µg), cefepime (30 µg), ceftazidime (30 µg), piperacillin/tazobactam (110 μg), ciprofloxacin (5 μg), tobramycin (10 μg), amikacin (30 μg), gentamicin (10 µg), ampicillin (10 µg), and aztreo-nam (30 µg). American composing gathering (ATCC 27853) of P. aeruginosa was utilized as a control strain to decide antibacterial powerlessness.

#### 4. RESULTS

Among 270 strains, 55 MBL-creating P. aeruginosa strains were confined from patients hospitalized in consume unit. Of the 55 consume cases, 29 (53%) were females and 26 (47%) guys. the age of the consume patients extended from 16 to 87 years, with most extreme number of cases in the age gathering of 16 to 36 years (n = 40; 72.7%) (Table 1). Fourteen consume harmed patients (25.5%) were from urban regions and 41 (74.5%) were from country. Thirty-three cases (60%) were unplanned, and 22 (40%) were self-destructive. By and large, 51 patients (93%) had concoction wounds, and 4 (7%) had electrical wounds. As opposed to guys, consumes because of synthetic ex-posures were progressively visit in females (52.7%). Of the 55 consume patients, 17 patients (30%) passed on. Females demonstrated a high frequency of mortality (females, 70.5%; guys, 29.5%; and female to male proportion, 2.4). All passings were because of concoction exposures (Table 2).

As per the in vitro anti-microbial powerlessness testing by E-test technique, ceftazidime was the best antibiotic. Thirty-five segregates (63.5%) were impervious to all the 11 tried anti-infection agents.

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Category	Results	
Age, y	35 ± 17	
16-36	40 (72.7)	
37-56	9 (16.3)	
>56	6 (11)	
Residence		
Rural	41 (74.5)	
Urban	14 (25.5)	
Hospitalization, d	$25 \pm 12$	
1-30	34 (61.8)	
31-60	15 (27.2)	
61-90	5 (9)	
91-120	1 (1)	

Table 2 Distribution Of The Burn Patients In Terms Of The Gender, Burn Cause, And Exposure Types

Gender	Results						
Male $(n = 26)$	8 urban						
	18 rural						
Female $(n = 29)$	6 urban						
	23 rural						
Burn cause							
Accidental $(n = 33)$	23 males						
	10 females						
Suicidal (n = 22)	3 males						
	19 females						
Exposure							
Electricity $(n = 4)$	4 females						
	0 male						
Chemical $(n = 51)$	29 female						
	22 males						
	29 females						

Table 3 Antibiotic Resistance Patterns Of Metallo-beta-lactamaseproducing *Pseudomonas Aeruginosa* Strains Isolated From Burn Patients

Antibiotic Resistance Patterns	Frequency(n = 55)
IMI, MEM, CAZ, ATM, TN, GM, PTZ, CPM,	35 (63.6)
AK,	
IMI, MEM, ATM, TN, GM, PTZ, CPM, AK,	8 (14.5)
CIP, AP	
IMI, MEM, ATM, TN, GM, PTZ, CPM, AK, AP	5 (9.1)
IMI, MEM, CAZ, ATM, TN, PTZ, CPM, AK,	1 (1.8)
CIP, AP	
MEM, CAZ, ATM, TN, GM, PTZ, CPM, AK,	1 (1.8)
CIP	
IMI, MEM, CAZ, ATM, PTZ, CPM, AP	1 (1.8)
IMI, MEM, CAZ, ATM, PTZ, CPM, AK, AP	1 (1.8)
IMI, CAZ, ATM, TN, GM, CPM, AK, AP	1 (1.8)
IMI, MEM, AK, AP	1 (1.8)
IMI, AK, CIP, AP	1 (1.8)

#### DISCUSSION

P. aeruginosa is an astute pathogen, which is normally connected with nosocomial contaminations (9). Antimicrobial obstruction amid treatment happens fre-quently among the at first vulnerable P. aeruginosa segregates, bringing about the development of protection from mul-tiple anti-infection agents (10). MBL-creating P. aeruginosa strains have been recognized from clinical separates worldwide with expanding recurrence in the course of recent years, and these disconnects are in charge of extended nosoco-mial diseases in various nations (11). The rising fre-quency of MDRPA strains involves worry as effec-tive antimicrobial choices are fundamentally restricted (12). In this examination, females were influenced more than guys (53% versus 47%). This might be because of increasingly visit contribution of females in family tasks, which request more ex-posure to flame (for example in cooking and warming). The outcomes demonstrated that the mortality in our consume cases was 30%, which is lower than the rates in some different reports (13) yet higher than the rates in a large portion of different nations (14). The all out body surface zone consumed (entire body in half and entire body with the exception of head in 41% of cases) was the most successive factor identified with mortality. The expanding sever-ity of wounds caused by more skin misfortune, uncovered the larg-est consumes to more complexities, in this way, rising death rate. Another noticeable finding in the present examination was the abnormal state of antibacterial obstruction profile identified in the secludes of P. aeruginosa. Protection from aztreonam, piperacillin/tazobactam, meropenem, imipenem, and ampicillin was found in all the disengages. Consequently, while regulating experimental treatment in consume patients with nosocomial diseases with P. aeruginosa, MBL ought to be considered on the off chance that the patients are not receptive to vehicle bapenem treatment. The dimension of MBL creation among MDRPA strains is by all accounts more prominent than that evaluated in Iran. Reports on MBLdelivering P. aeruginosa segregates are expanding all inclusive because of the expanded beta-lactam us-age and rise of safe microscopic organisms under anti-infection weight. As of now, CLSI archive has no rules for distinguishing MBLs in P. aeruginosa. This disturbing antibiot-ic obstruction slant was seen for P. aeruginosa strains, as observed in past investigations (15). A comparable report of MDRPA was additionally detailed by different specialists. In spite of the fact that ceftazi-dime was observed to be the best medication in the pres-ent consider, protection from this anti-toxin was high (76.6%). In one examination from Iran, higher protection from ceftazidime was accounted for (16); this could be because of the reason that these are save sedates and are utilized if all else fails for MDRPA in healing facility consume settings in Shiraz. In the cur-rent examine, the microscopic organisms segregated from just nine patients getting experimental anti-toxin treatment were delicate to the endorsed antimicrobials as indicated by MIC technique. The adjustments in the bacterial opposition designs, as saw in the consume wards, could have essential ramifications for both clinical settings and epidemiological purposes. Such a high anti-toxin obstruction in P. aeruginosa disconnects is likely because of the particular weight applied on the microorganisms in view of the elements, for example, poor adherence to healing facility anti-microbial arrangement and over the top just as indis-criminate utilization of wide range antimicrobial specialists including beta-lactams, carbapenems, aminoglycosides, and quinolones. These MDRPA strains set up them-selves in the clinic

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condition and, accordingly, spread starting with one patient then onto the next, from therapeutic faculty to the patients, or among various units in the healing facility. All the more as of late, MDRPA and pandrug-safe P. aeruginosa detaches have risen in healing center consume units. Henceforth, there is not really any powerful anti-toxin against pandrug-safe strains, in which an external film boundary of low porousness and a variety of effective multidrug efflux siphons are joined with large numbers of explicit anti-microbial obstruction systems. Treatment of the in-fections caused by these supposed "superbugs" stays testing in light of the fact that the pool of successful anti-infection agents is contracting and couple of new antipseudomonal anti-infection agents are being developed. In consume patients, a successful and con-tinuous reconnaissance for disease control and accordingly, customary tissue refined for control purposes no less than two times every week are prescribed. Routine microbiologi-cal reconnaissance and watchful in vitro testing of antiinfection agents before solution and strict adherence to the healing facility anti-toxin arrangement might be useful in the counteractive action, treat-ment, and control of MDRPA in the patients hospitalized in consume units. Further examinations ought to be done to decide the changing affectability profiles of the dif-ferent gram-positive and Gram-negative microbes and individual antimicrobial vulnerability designs (17-20). In such manner, the doctor's facilities ought to figure an effec-tive anti-toxin approach. At the point when the injury bacterial checks are in excess of 105 microorganisms for each gram of tissue, the danger of wound disease is extraordinary, skin unite survival is poor, and wound mending is deferred (21). Bacterial tallies of under 103 life forms for each gram, are not typically inva-sive and permit skin unite survival rates of over 90%. The objectives of the neighborhood wound administration ought to be the counteractive action of practical tissues drying up and the control of microscopic organisms.

### REFERENCES

- Laupland KB, Parkins MD, Church DL, et al. Population-based epidemiological study f infections caused by carbapenem-resistant Pseudomonas aeruginosa in the Calgary Health Region: Importance of metallo- β-lactamase (MBL)-producing strains. J Infect Dis. 2005;192:1606-1612.
- 2 Bush K, Jacoby GA, Medeiros AA, A functional classification scheme for B-lactamases correlation with molecular structure. Antimicrob Agents Chemother. 1995:39:1211-1233.
- 3. Livermore DM, Woodford N. Carbapenemases: A problem in waiting? Curr Opin Microbiol. 2000;3:489-495.
- 4. Nordmann P, Poirel L. Emerging carbapenemases in gram negative aerobes. Clin Microbiol Infect. 2002;8:321-331
- Sacha P, Wieczorek P, Hauschild T, et al. Metallo-beta-lactamases of Pseudomonas 5. aeruginosa\_ -A novel mechanism resistance to β-lactam antibiotics. Folia Histochem Cytobiol. 2008;46:137-142
- Pitout JDD, Gregson DB, Poirel L, et al. Detection of Pseudomonas aeruginosa 6 producing metallo-beta-lactamases in a large centralized laboratory. J Clin Microbiol. 2005;43:3129-3135.
- 7 Khosravi AD, Mihani F. Detection of metallo-beta-lactamase-producing Pseudomonas aeruginosa strains isolated from burn patients in Ahwaz, Iran. Diagn Microbiol Infect Dis. 2008:60:125-128.
- Japoni A, Alborzi A, Kalani M, et al. Susceptibility patterns and crossresistance of 8. artibiotics against Pseudomonas aeruginosa isolated from burn patients in the South of Iran. Burns. 2006;32:343-347.
- Sharma BR, Singh VP, Bangar S, Gupta N. Septicemia: The Principal Killer of Burns Patients. American Journal of Infectious Diseases. 2005;1(3):132–38. 9
- Ho SA, Lee TW, Denton M, Conway SP, Brownlee KG. Regimens for eradicating early Pseudomonas aeruginosa infection in children do not promote antibiotic resistance in 10
- Data T, Huang YY, Sharma SK, Hashmi JT, Kurup DB, Hamblin MR. Topical antimicrobials for burn wound infections. Recent Pat Antiinfect Drug Discov. 11. 2010;5(2):124-51.
- Japoni A, Farshad S, Alborzi A. Pseudomonas aeruginosa: Burn Infection, Treatment and Antibacterial Resistance. IRCMJ. 2009;11(3). Mayhall CG. The epidemiology of burn wound infections: then and now. Clin Infect Dis. 12.
- 13. 2003:37(4):543-50
- 2005,37(4).343-350 McVay CS, Velasquez M, Fralick JA. Phage therapy of Pseudomonas aeruginosa infection in a mouse burn wound model. Antimicrob Agents Chemother. 2007;51(6):1934-8. 14.
- Pirs M, Andlovic A, Cerar T, Zohar-C'retnik T, Kobola L, Kolman J, et al. A case of 15 FIRS M, Andrew R, Cotta F, Evan F, Frank F, Henne H, H
- Cardoso T, Ribeiro O, Aragao IC, Costa-Pereira A, Sarmento AE. Additional risk factors for infection by multidrug-resistant pathogens in healthcare-associated infection: a large 16. cohort study. BMC Infect Dis. 2012;12:375. Mardaneh J, Ahmadi K, Jahan Sepas A. Determination antimicrobial resistance profile
- of Pseudomonas aeruginosa strains isolated from hospitalized patients in Taleghani Hospital (Ahvaz, Iran) from 2011-2012. JFUMS. 2013;3(3):188-93.
- Pagani L, Mantengoli E, Migliavacca R, Nucleo E, Pollini S, Spalla M, et al. Multifocal detection of multidrug-resistant Pseudomonas aeruginosa producing the PER-1 18 extended-spectrum beta-lactamase in Northern Italy. J Clin Microbiol. 2004;42(6):2523-9.
- Obritsch MD, Fish DN, MacLaren R, Jung R. Nosocomial infections due to multidrug-19 resistant Pseudomonas aeruginosa: epidemiology and treatment options. Pharmacotherapy. 2005;25(10):1353-64.
- Al-Ibran E. Rao MH. Fatima K. Irfan S. Johal M. Khan M. Current Bacteriological 20 profile in Fire-burn victims and their associated mortality at the Burns Centre, Karachi-Pakistan, Pakistan Journal of Medical Sciences, 2011:27(4).
- 21. Keen EF, 3rd, Robinson BJ, Hospenthal DR, Aldous WK, Wolf SE, Chung KK, et al. Incidence and bacteriology of burn infections at a military burn center. Burns. 2010;36(4):461-8.

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